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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants: Iva Greenwald and Diane Levitan

Serial No.: 09/043,944 Examiner: Samuel Wei Liu

Filed: March 27, 1998 Group Art Unit: 1653

For: IDENTIFICATION OF SEL-12 AND USES THEREOF

1185 Avenue of the Americas  
New York, NY 10036  
February 13, 2003

Assistant Commissioner for Patents  
Washington, D.C. 20231

Sir:

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT

In accordance with their duty of disclosure under 37 C.F.R. §1.56, applicants request that the following disclosures be made of record in the above-identified application pursuant to 37 C.F.R. §1.97(c). These items are also listed on the attached Form PTO-1449 (**Exhibit A**). Copies of these items are attached hereto as **Exhibits 1-24** respectively.

1. PCT International Application No. WO 97/11956, published April 3, 1997 (**Exhibit 1**);
2. European Search Report, dated November 20, 2002 (**Exhibit 2**);
3. PCT International Search Report, dated January 21, 1997 (**Exhibit 3**);

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4. PCT Written Opinion, dated July 15, 1997  
**(Exhibit 4);**
5. Data Base Search, total of 3 pages **(Exhibit 5);**
6. Bai, C. et al., "SKP1 connects cell cycle regulators to the ubiquitin proteolysis machinery through a novel motif, the F-box," Cell 86:263-74 (1996)  
**(Exhibit 6);**
7. Ellisen, L.W. et al., "TAN-1, the human homolog of the Drosophila Notch gene, is broken by chromosomal translocations in T lymphoblastic neoplasms," Cell 66:649-61 (1991) **(Exhibit 7);**
8. Gallahan, D. and Callahan, R., "Mammary tumorigenesis in feral mice: identification of a new int locus in mouse mammary tumor virus (Czech II)-induced mammary tumors," J. Virol. 61:66-74 (1987) **(Exhibit 8);**
9. Grant, B. and Greenwald, I., "Structure, function and expression of SEL-1, a negative regulator of LIN-2 and GLP-1 in C. elegans," Development 124:637-644 (1997) **(Exhibit 9);**
10. Greenwald, I. et al., "The lin-12 locus specifies cell fates in C. elegans," Cell 34:435-44 (1983)  
**(Exhibit 10);**

11. Greenwald, I. and Seydoux, G., "Analysis of gain-of-function mutations of the lin-2 gene of *C. elegans*," *Nature* 346:197-99 (1990) (**Exhibit 11**);
12. Hubbard, J. et al., "Sel-10 negative regulator of lin-12 activity in *C. elegans*, encodes a member of the CDC4 family of proteins," *Genes Dev.* 11:3182-93 (1997) (**Exhibit 12**);
13. Kimble, J., "Alteration in cell lineage following laser ablation of cells in the somatic gonad of *C. elegans*," *Dev. Biol.* 87:286-300 (1981) (**Exhibit 13**);
14. King, R.W. et al, "How proteolysis drives the cell cycle," *Science* 274:1652-58 (1996) (**Exhibit 14**);
15. Neer, E.J. et al., "The ancient regulatory-protein family of WD-repeat proteins," *Nature* 371:297-300 (1994) (**Exhibit 15**);
16. Robbins, J. et al. "Mouse mammary tumor gene int-3: a member of the Notch gene family transforms mammary epithelial cells," *J. Virol.* 66:2594-99 (1992) (**Exhibit 16**);
17. Seydoux, G. and Greenwald, I., "Cell autonomy of lin-12 function in a cell fate decision in *C. elegans*," *Cell* 57:1237-45 (1989) (**Exhibit 17**);

18. Shen, J. et al, "Skeletal and CNS defects in presenilin-1-deficient mice," *Cell* 89:629-39 (1997) **(Exhibit 18)**;
19. Sherrington, R. et al., "Cloning of a gene bearing missense mutations in early-onset familial Alzheimer's disease," *Nature* 375:754-60 (1995) **(Exhibit 19)**;
20. Sundaram, M. and Greenwald, I., "Suppressors of a lin-12 hypomorph define genes that interact with both lin-12 and glp-1 in *C. elegans*," *Genetics* 135: 765-83 (1993) **(Exhibit 20)**;
21. Tuck, S. and Greenwald, I., "Lin-25, a gene required for vulval induction in *C. elegans*," *Genes Dev.* 9:341-57 (1995) **(Exhibit 21)**;
22. Wilson, R. et al., "2.2 Mb of contiguous nucleotide sequence from chromosome III of *C. elegans*," *Nature* 368:32-38 (1994) **(Exhibit 22)**;
23. Wong, P.C. et al., "Presenilin-1 is required for Notch1 and DII 1 expression in the paraxial mesoderm," *Nature* 387:288-91 (1997) **(Exhibit 23)**;  
and

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24. Yochem, J. and Byers, B., "Structural comparison of the yeast cell division cycle gene CDC4 and a related pseudogene," J. Mol. Biol. 195:233-45 (1987) **(Exhibit 24)**.

On November 20, 2002, a Search Report was issued in connection with European Patent Application No. 98935974, based on PCT International Application PCT/US98/15335, which is related to the subject application, citing items 1, 12, and 20 above. A copy of the November 20, 2002 Search Report is attached hereto as **Exhibit 2**. Item 5 above was cited in a PCT Written Opinion issued in connection with PCT/US98/15335. A copy of the PCT Written Opinion is attached hereto as **Exhibit 4**.

No fee, other than the enclosed \$180.00 fee, is deemed necessary in connection with the filing of this Information Disclosure Statement. However, if any additional fee is required, authorization is hereby given to charge the amount of any such fee to Deposit Account No. 03-3125.

Respectfully submitted,

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I hereby certify that this correspondence is being deposited this date with the U.S. Postal Service with sufficient postage as first class mail in an envelope addressed to:  
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